## FIG. 1A

|   |   |   |      |   |   |   |   |  |  |   | atc<br>Ile<br>15  | 48  |
|---|---|---|------|---|---|---|---|--|--|---|-------------------|-----|
| _ | _ | _ | <br> | _ | - | _ | _ |  |  | _ | agc<br>Ser        | 96  |
|   |   |   |      |   |   |   |   |  |  |   | cgc<br>Arg        | 144 |
|   |   |   |      |   |   |   |   |  |  |   | gtc<br>Val        | 192 |
|   |   |   |      |   |   |   |   |  |  |   | gcg<br>Ala        | 240 |
|   |   |   |      |   |   |   |   |  |  |   | ctg<br>Leu<br>95  | 288 |
|   |   |   |      |   |   |   |   |  |  |   | gac<br>Asp        | 336 |
|   |   |   |      |   |   |   |   |  |  |   | tgg<br>Trp        | 384 |
|   |   |   |      |   |   |   |   |  |  |   | atg<br>Met        | 432 |
|   |   |   |      |   |   |   |   |  |  |   | aaa<br>Lys        | 480 |
|   |   |   |      |   |   |   |   |  |  |   | gag<br>Glu<br>175 | 528 |
|   |   |   |      |   |   |   |   |  |  |   | atg<br>Met        | 576 |
|   |   |   |      |   |   |   |   |  |  |   | ttc<br>Phe        | 624 |
|   |   |   |      |   |   |   |   |  |  |   | tat<br>Tyr        | 672 |

## FIG. 1B

|            |                   |            |                   |                   | tca<br>Ser<br>230 |                   |            |                   |                   |            |                   |            |                   |                   |            | 720  |
|------------|-------------------|------------|-------------------|-------------------|-------------------|-------------------|------------|-------------------|-------------------|------------|-------------------|------------|-------------------|-------------------|------------|------|
|            |                   |            |                   |                   | tgt<br>Cys        |                   |            |                   |                   |            |                   |            |                   |                   |            | 768  |
| ata<br>Ile | agg<br>Arg        | tcg<br>Ser | ctc<br>Leu<br>260 | aca<br>Thr        | gag<br>Glu        | cgg<br>Arg        | ctc<br>Leu | tat<br>Tyr<br>265 | gtc<br>Val        | ggg<br>Gly | ggt<br>Gly        | ccc<br>Pro | atg<br>Met<br>270 | act<br>Thr        | aac<br>Asn | 816  |
|            |                   |            |                   |                   | tgc<br>Cys        |                   |            |                   |                   |            |                   |            |                   |                   |            | 864  |
| _          | _                 |            | _                 | _                 | ggt<br>Gly        |                   |            |                   |                   | _          |                   | _          | _                 | _                 | _          | 912  |
|            |                   |            |                   |                   | gcc<br>Ala<br>310 |                   |            |                   |                   |            |                   |            |                   |                   |            | 960  |
| gga<br>Gly | gac<br>Asp        | gac<br>Asp | ctt<br>Leu        | gtc<br>Val<br>325 | gtt<br>Val        | atc<br>Ile        | tgt<br>Cys | gaa<br>Glu        | agc<br>Ser<br>330 | gcg<br>Ala | gga<br>Gly        | acc<br>Thr | caa<br>Gln        | gag<br>Glu<br>335 | gac<br>Asp | 1008 |
| gcg<br>Ala | gca<br>Ala        | agc<br>Ser | cta<br>Leu<br>340 | cga<br>Arg        | gtc<br>Val        | ttc<br>Phe        | acg<br>Thr | gag<br>Glu<br>345 | gct<br>Ala        | atg<br>Met | act<br>Thr        | agg<br>Arg | tac<br>Tyr<br>350 | tct<br>Ser        | gcc<br>Ala | 1056 |
|            |                   |            |                   |                   | ccc<br>Pro        |                   |            |                   |                   |            |                   |            |                   |                   |            | 1104 |
| tcg<br>Ser | tgt<br>Cys<br>370 | tcc<br>Ser | tcc<br>Ser        | aat<br>Asn        | gtg<br>Val        | tcg<br>Ser<br>375 | gtc<br>Val | gca<br>Ala        | cac<br>His        | gat<br>Asp | gca<br>Ala<br>380 | tct<br>Ser | ggc<br>Gly        | aaa<br>Lys        | agg<br>Arg | 1152 |
|            |                   |            |                   |                   | cgt<br>Arg<br>390 |                   |            |                   |                   |            |                   |            |                   |                   |            | 1200 |
|            |                   |            |                   |                   | cac<br>His        |                   |            |                   |                   |            |                   |            |                   |                   |            | 1248 |
|            |                   |            |                   |                   | act<br>Thr        |                   |            |                   |                   |            |                   |            |                   |                   |            | 1296 |
|            |                   |            |                   |                   | cta<br>Leu        |                   |            |                   |                   |            |                   |            |                   |                   |            | 1344 |

# FIG. 1C

|            |            |            |                   |            |            |            |            | tcc<br>Ser        |            |            |            |            |                   |            |            | 1392 |
|------------|------------|------------|-------------------|------------|------------|------------|------------|-------------------|------------|------------|------------|------------|-------------------|------------|------------|------|
|            |            |            |                   |            |            |            |            | ctt<br>Leu        |            |            |            |            |                   |            |            | 1440 |
|            |            |            |                   |            |            |            |            | gtg<br>Val        |            |            |            |            |                   |            |            | 1488 |
|            |            |            |                   |            |            |            |            | aga<br>Arg<br>505 |            |            |            |            |                   |            |            | 1536 |
|            |            |            |                   |            |            |            |            | agg<br>Arg        |            |            |            |            |                   |            |            | 1584 |
|            |            |            |                   |            |            |            |            | aag<br>Lys        |            |            |            |            |                   |            |            | 1632 |
|            |            |            |                   |            |            |            |            | ggc<br>Gly        |            |            |            |            |                   |            |            | 1680 |
|            |            |            |                   |            |            |            |            | tct<br>Ser        |            |            |            |            |                   |            |            | 1728 |
| atg<br>Met | ttg<br>Leu | tgc<br>Cys | cta<br>Leu<br>580 | ctc<br>Leu | cta<br>Leu | ctc<br>Leu | tcc<br>Ser | gta<br>Val<br>585 | gly<br>aaa | gta<br>Val | ggc<br>Gly | atc<br>Ile | tat<br>Tyr<br>590 | cta<br>Leu | ctc<br>Leu | 1776 |
|            |            |            |                   | cac<br>His |            |            |            |                   |            |            |            |            |                   |            |            | 1803 |

## FIG. 2A

| atg<br>Met<br>1   | gct<br>Ala        | agc<br>Ser        | atg<br>Met        | tca<br>Ser<br>5   | atg<br>Met        | tcc<br>Ser        | tat<br>Tyr        | aca<br>Thr        | tgg<br>Trp<br>10  | aca<br>Thr        | ggc<br>Gly        | gcc<br>Ala        | ctg<br>Leu        | atc<br>Ile<br>15  | aca<br>Thr        | 48  |
|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-----|
|                   |                   |                   |                   |                   |                   | agc<br>Ser        |                   |                   |                   |                   |                   |                   |                   |                   |                   | 96  |
|                   |                   |                   |                   |                   |                   | aac<br>Asn        |                   |                   |                   |                   |                   |                   |                   |                   |                   | 144 |
| gca<br>Ala        | agc<br>Ser<br>50  | ctg<br>Leu        | cgg<br>Arg        | cag<br>Gln        | aag<br>Lys        | aag<br>Lys<br>55  | gtc<br>Val        | acc<br>Thr        | ttt<br>Phe        | gac<br>Asp        | aga<br>Arg<br>60  | ctg<br>Leu        | caa<br>Gln        | gtc<br>Val        | ctg<br>Leu        | 192 |
|                   |                   |                   |                   |                   |                   | gtg<br>Val        |                   |                   |                   |                   |                   |                   |                   |                   |                   | 240 |
|                   |                   |                   |                   |                   |                   | cta<br>Leu        |                   |                   |                   |                   |                   |                   |                   |                   |                   | 288 |
| ccc<br>Pro        | ccg<br>Pro        | cat<br>His        | tcg<br>Ser<br>100 | gcc<br>Ala        | aaa<br>Lys        | tcc<br>Ser        | aaa<br>Lys        | ttt<br>Phe<br>105 | ggc               | tat<br>Tyr        | gly<br>aaa        | gca<br>Ala        | aag<br>Lys<br>110 | gac<br>Asp        | gtc<br>Val        | 336 |
|                   |                   |                   |                   |                   |                   | gcc<br>Ala        |                   |                   |                   |                   |                   |                   |                   |                   |                   | 384 |
| gac<br>Asp        | ttg<br>Leu<br>130 | ctg<br>Leu        | gag<br>Glu        | gac<br>Asp        | act<br>Thr        | gac<br>Asp<br>135 | aca<br>Thr        | cca<br>Pro        | att<br>Ile        | cag<br>Gln        | acc<br>Thr<br>140 | acc<br>Thr        | atc<br>Ile        | atg<br>Met        | gca<br>Ala        | 432 |
| aaa<br>Lys<br>145 | aat<br>Asn        | gag<br>Glu        | gtt<br>Val        | ttc<br>Phe        | tgc<br>Cys<br>150 | gtc<br>Val        | caa<br>Gln        | cca<br>Pro        | gag<br>Glu        | aaa<br>Lys<br>155 | gga<br>Gly        | ggc<br>Gly        | cgc<br>Arg        | aaa<br>Lys        | cca<br>Pro<br>160 | 480 |
| gct<br>Ala        | cgc<br>Arg        | ctc<br>Leu        | atc<br>Ile        | gta<br>Val<br>165 | ttc<br>Phe        | cca<br>Pro        | gac<br>Asp        | ctg<br>Leu        | gga<br>Gly<br>170 | gtt<br>Val        | cgt<br>Arg        | gta<br>Val        | tgc<br>Cys        | gag<br>Glu<br>175 | aag<br>Lys        | 528 |
| atg<br>Met        | gcc<br>Ala        | ctt<br>Leu        | tac<br>Tyr<br>180 | gac<br>Asp        | gtg<br>Val        | gtt<br>Val        | tcc<br>Ser        | act<br>Thr<br>185 | ctt<br>Leu        | cct<br>Pro        | cag<br>Gln        | gcc<br>Ala        | gtg<br>Val<br>190 | atg<br>Met        | ggc<br>Gly        | 576 |
| tcc<br>Ser        | tca<br>Ser        | tac<br>Tyr<br>195 | gga<br>Gly        | ttc<br>Phe        | caa<br>Gln        | tac<br>Tyr        | tct<br>Ser<br>200 | cct<br>Pro        | aag<br>Lys        | cag<br>Gln        | cgg<br>Arg        | gtc<br>Val<br>205 | gag<br>Glu        | ttc<br>Phe        | ctg<br>Leu        | 624 |
| gtg<br>Val        | aat<br>Asn<br>210 | acc<br>Thr        | tgg<br>Trp        | aaa<br>Lys        | gca<br>Ala        | aag<br>Lys<br>215 | aaa<br>Lys        | tgc<br>Cys        | cct<br>Pro        | atg<br>Met        | ggc<br>Gly<br>220 | ttc<br>Phe        | tca<br>Ser        | tat<br>Tyr        | gac<br>Asp        | 672 |

### FIG. 2B

| acc<br>Thr<br>225 | cgc<br>Arg | tgt<br>Cys        | ttt<br>Phe | gac<br>Asp        | tca<br>Ser<br>230 | acg<br>Thr | gtc<br>Val        | act<br>Thr | gag<br>Glu        | aat<br>Asn<br>235 | gac<br>Asp | atc<br>Ile        | cgt<br>Arg | gtt<br>Val        | gag<br>Glu<br>240 | 720  |
|-------------------|------------|-------------------|------------|-------------------|-------------------|------------|-------------------|------------|-------------------|-------------------|------------|-------------------|------------|-------------------|-------------------|------|
|                   |            |                   |            |                   | tgt<br>Cys        |            |                   |            |                   |                   | _          | _                 | _          | _                 | _                 | 768  |
|                   |            |                   |            |                   | gag<br>Glu        |            |                   |            |                   |                   |            |                   |            |                   |                   | 816  |
|                   |            |                   |            |                   | tgc<br>Cys        |            |                   |            |                   |                   |            |                   |            |                   |                   | 864  |
|                   |            |                   |            |                   | ggt<br>Gly        |            |                   |            |                   |                   |            |                   |            |                   |                   | 912  |
| gca<br>Ala<br>305 | gcc<br>Ala | tgt<br>Cys        | cga<br>Arg | gct<br>Ala        | gcc<br>Ala<br>310 | aag<br>Lys | ctc<br>Leu        | cag<br>Gln | gac<br>Asp        | tgc<br>Cys<br>315 | acg<br>Thr | atg<br>Met        | ctc<br>Leu | gtg<br>Val        | aat<br>Asn<br>320 | 960  |
|                   |            |                   |            |                   | gtt<br>Val        |            |                   |            |                   |                   |            |                   |            |                   |                   | 1008 |
|                   |            |                   |            |                   | gtc<br>Val        |            |                   |            |                   |                   |            |                   |            |                   |                   | 1056 |
|                   |            |                   |            |                   | ccc<br>Pro        |            |                   |            |                   |                   |            |                   |            |                   |                   | 1104 |
|                   |            |                   |            |                   | gtg<br>Val        |            |                   |            |                   |                   |            |                   |            |                   |                   | 1152 |
| gtg<br>Val<br>385 | tac<br>Tyr | tac<br>Tyr        | ctc<br>Leu | acc<br>Thr        | cgt<br>Arg<br>390 | gac<br>Asp | ccc<br>Pro        | acc<br>Thr | gtc<br>Val        | ccc<br>Pro<br>395 | ctt<br>Leu | gcg<br>Ala        | cgg<br>Arg | gct<br>Ala        | gcg<br>Ala<br>400 | 1200 |
| tgg<br>Trp        | gag<br>Glu | aca<br>Thr        | gct<br>Ala | agg<br>Arg<br>405 | cac<br>His        | act<br>Thr | cca<br>Pro        | gtc<br>Val | aac<br>Asn<br>410 | tcc<br>Ser        | tgg<br>Trp | cta<br>Leu        | ggc<br>Gly | aac<br>Asn<br>415 | atc<br>Ile        | 1248 |
|                   |            |                   |            |                   | act<br>Thr        |            |                   |            |                   |                   |            |                   |            |                   |                   | 1296 |
| ttc<br>Phe        | ttc<br>Phe | tcc<br>Ser<br>435 | atc<br>Ile | ctt<br>Leu        | cta<br>Leu        | gcc<br>Ala | cag<br>Gln<br>440 | gag<br>Glu | caa<br>Gln        | ctt<br>Leu        | gaa<br>Glu | aaa<br>Lys<br>445 | gcc<br>Ala | ctg<br>Leu        | gat<br>Asp        | 1344 |

## FIG. 2C

|   |            |  | Gly<br>999        |   |   |  |  |   |   |  |   | -        | 1392 |
|---|------------|--|-------------------|---|---|--|--|---|---|--|---|----------|------|
|   |            |  | cga<br>Arg        |   |   |  |  |   |   |  |   | -        | 1440 |
|   |            |  | gag<br>Glu<br>485 |   |   |  |  |   |   |  |   | =        | 1488 |
|   |            |  | ttg<br>Leu        |   |   |  |  |   |   |  |   | -        | 1536 |
|   |            |  | tcc<br>Ser        |   |   |  |  |   |   |  |   | -        | 1584 |
|   |            |  | gca<br>Ala        |   |   |  |  |   |   |  |   | <u>-</u> | 1632 |
| _ |            |  | ctg<br>Leu        | _ | _ |  |  | _ | _ |  | _ | -        | 1680 |
|   |            |  | tat<br>Tyr<br>565 |   |   |  |  |   |   |  |   | -        | 1728 |
|   | cac<br>His |  |                   |   |   |  |  |   |   |  |   | 3        | 1740 |

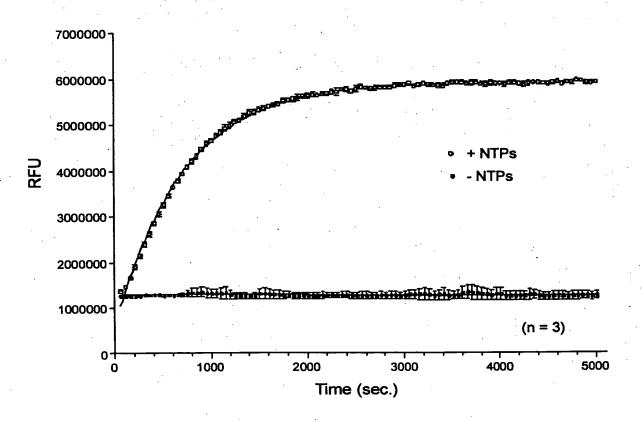


FIG. 4

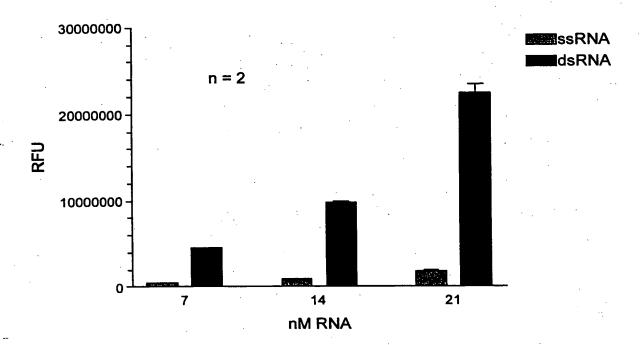


FIG. 5A

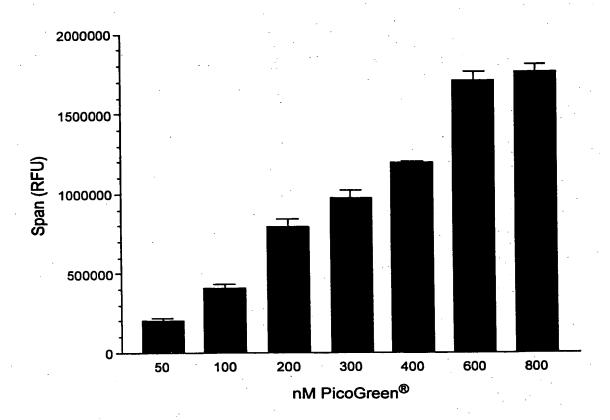


FIG. 5B

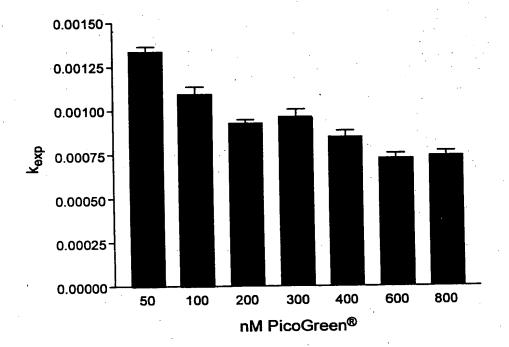


FIG. 5C

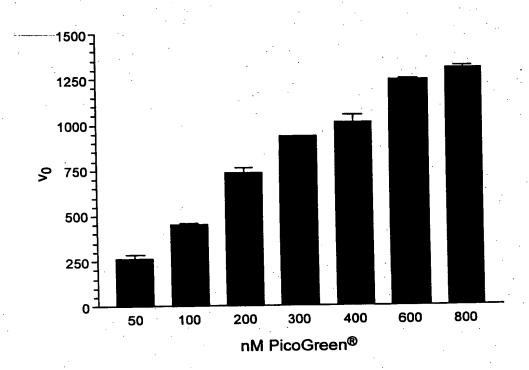


FIG. 6

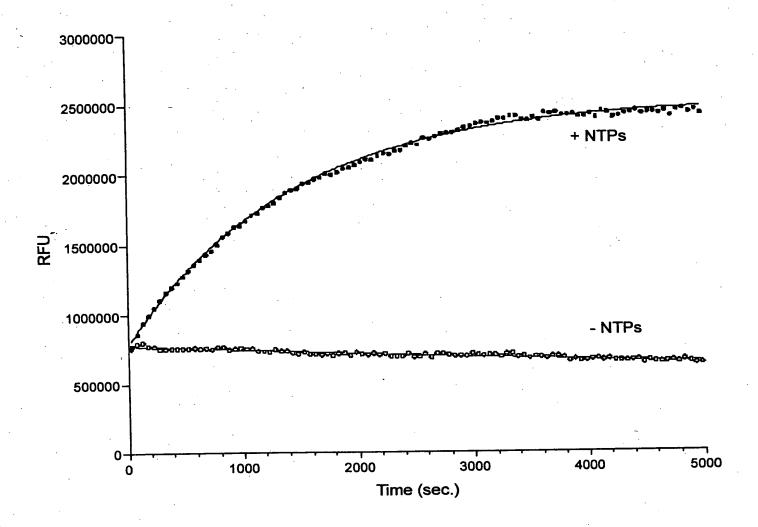


FIG. 7

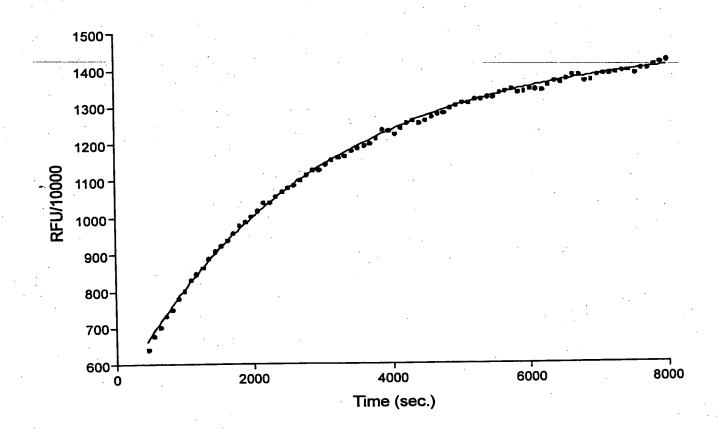


FIG. 8

